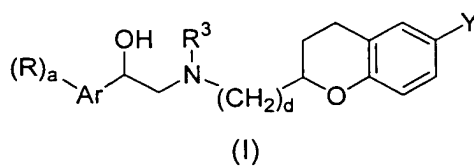




We claim:

1. A compound of Formula I



wherein,-

R is independently

- 10
- hydroxy,
  - oxo,
  - halo,
  - cyano,
  - nitro,
  - C<sub>1</sub>-C<sub>10</sub> alkyl,
  - 15 • C<sub>1</sub>-C<sub>10</sub> haloalkyl,
  - CF<sub>3</sub>,
  - NR<sup>1</sup>R<sup>1</sup>,
  - SR<sup>1</sup>,
  - OR<sup>1</sup>,
  - 20 • SO<sub>2</sub>R<sup>2</sup>,
  - OCOR<sup>2</sup>,
  - NR<sup>1</sup>COR<sup>2</sup>,
  - COR<sup>2</sup>,
  - NR<sup>1</sup>SO<sub>2</sub>R<sup>2</sup>,
  - 25 • phenyl, or
  - a 5- or 6-membered heterocycle with from 1 to 4 heteroatoms selected from O, S, and N;
- each cyclic moiety being optionally substituted with
- 30
- hydroxy,
  - R<sup>1</sup>,
  - halo,
  - cyano,

- $\text{NR}^1\text{R}^1$ ,
- $\text{SR}^1$ ,
- $\text{CF}_3$ ,
- $\text{OR}^1$ ,
- 5      •  $\text{C}_3\text{-C}_8$  cycloalkyl,
- $\text{NR}^1\text{COR}^2$ ,
- $\text{COR}^2$ ,
- $\text{SO}_2\text{R}^2$ ,
- $\text{OCOR}^2$ ,
- 10      •  $\text{NR}^1\text{SO}_2\text{R}^2$ ,
- $\text{C}_1\text{-C}_{10}$  alkyl, or
- $\text{C}_1\text{-C}_{10}$  alkoxy;

$\text{R}^1$  is

- 15      • hydrogen,
  - $(\text{CH}_2)_d\text{-O-(CH}_2)_d\text{R}^5$  where each  $d$  is selected independently, or
  - $\text{C}_1\text{-C}_{10}$  alkyl optionally substituted with 1 to 4 substituents each independently selected from
    - hydroxy,
    - 20      • halo,
    - $\text{CO}_2\text{C}_1\text{-C}_4\text{-alkyl}$ ,
    - $\text{CO}_2\text{H}$ ,
    - $\text{C}_1\text{-C}_{10}$  alkoxy,
    - $\text{S(O)}_b\text{C}_1\text{-C}_{10}$  alkyl,
    - 25      •  $\text{S(O)}_b\text{-phenyl}$  optionally substituted with halo,  $\text{C}_1\text{-C}_4$  alkyl,  $\text{C}_1\text{-C}_4$  alkoxy,  $\text{SO}_2\text{-C}_1\text{-C}_4\text{alkyl}$ , or  $\text{CO}_2\text{ C}_1\text{-C}_4\text{alkyl}$ ; or
    - phenyl optionally substituted with  $\text{CO}_2\text{C}_1\text{-C}_4\text{-alkyl}$ ,  $\text{CO}_2\text{H}$ , halo, or  $\text{C}_1\text{-C}_{10}$  alkyl;
  - or
  - 30      •  $\text{C}_3\text{-C}_8$  cycloalkyl, phenyl, or naphthyl, each optionally substituted with 1 to 4 substituents each independently selected from halo, nitro, oxo,  $\text{C}_1\text{-C}_{10}$  alkyl,  $\text{C}_1\text{-C}_{10}$  alkoxy,  $\text{C}_1\text{-C}_{10}$  alkylthio,  $\text{CO}_2\text{C}_1\text{-C}_4\text{-alkyl}$ , and  $\text{CO}_2\text{H}$ ,
- and

when two  $\text{R}^1$  groups are attached to N as  $\text{NR}^1\text{R}^1$ , these  $\text{R}^1$  groups may form together with the nitrogen to which they are attached, a heterocyclic ring

containing 4 to 7 C atoms, 1 to 2 N atoms, and 0 to 1 O or S atoms;

R<sup>2</sup> is

- R<sup>1</sup>,
  - OR<sup>1</sup>,
  - NR<sup>1</sup>R<sup>1</sup>,
  - NHS(O)<sub>b</sub>phenyl optionally substituted with C<sub>1</sub>-C<sub>4</sub> alkyl, C<sub>1</sub>-C<sub>4</sub> alkoxy, halo or nitro;
  - NHS(O)<sub>b</sub>naphthyl,
  - NHS(O)<sub>b</sub>C<sub>1</sub>-C<sub>10</sub> alkyl optionally substituted with fluoro up to the perfluoro level,
- or
- a 5- or 6-membered heterocycle with one or more heteroatoms selected from O, S, and N, said heterocyclic moiety being optionally substituted with R<sup>1</sup>;

R<sup>3</sup> is hydrogen, C<sub>1</sub>-C<sub>10</sub> alkyl, or COR<sup>2</sup>;

R<sup>4</sup> is hydrogen, C<sub>1</sub>-C<sub>10</sub> alkyl, C<sub>1</sub>-C<sub>10</sub> alkyl-phenyl, or C<sub>1</sub>-C<sub>10</sub> alkyl-pyridyl;

R<sup>5</sup> is hydrogen or COOH;

R<sup>6</sup> is

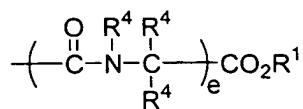
- hydrogen,
- C<sub>1</sub>-C<sub>10</sub> alkyl optionally substituted with 1 to 4 substituents each independently selected from halo, phenyl, or phenyl-COR<sup>2</sup>, or
- C<sub>1</sub>-C<sub>10</sub> alkyl-S(O)<sub>b</sub>C<sub>1</sub>-C<sub>10</sub> alkyl optionally substituted with COR<sup>2</sup> or C<sub>3</sub>-C<sub>8</sub> cycloalkyl;

Ar is

- phenyl optionally fused to a 5- or 6-membered heterocycle containing one or more heteroatoms each independently selected from O, S, and N, said bicyclic moiety being optionally fused to a phenyl, or
- a 5- or 6-membered heterocycle containing one or more heteroatoms each independently selected from N, S, and O, optionally fused to phenyl;

Y is

- halo,
- NO<sub>2</sub>,
- R<sup>6</sup>,
- SR<sup>1</sup>,
- S(O)<sub>b</sub>-phenyl-CO<sub>2</sub>R<sup>1</sup>,



where, when the two R<sup>4</sup> groups attached to the same C are both alkyl, they optionally may be joined so that, when taken together with the C to which they are attached, they form a spiro ring of 3, 5, or 6 C atoms, or where the R<sup>4</sup> attached to N and one R<sup>4</sup> attached to the adjacent C are both alkyl, they optionally may be joined so that, taken together with the atoms to which they are attached, they form a 5- or 6-membered heterocyclic ring;

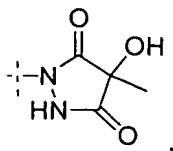
with the proviso that when e is 1, at least one R<sup>4</sup> group must be C<sub>1</sub>-C<sub>10</sub> alkyl-phenyl or C<sub>1</sub>-C<sub>10</sub> alkyl-pyridyl, or two R<sup>4</sup> groups must form one of said spiro or heterocyclic ring moieties;

- phenyl optionally fused to one or two phenyl rings, or to a 5- or 6-membered heterocycle containing one or more heteroatoms each independently selected from N, S, and O, or
  - a 5- or 6-membered heterocycle containing one or more heteroatoms each independently selected from N, S and O, optionally fused to a phenyl ring,
- each cyclic moiety being optionally substituted with one or more substituents independently selected from

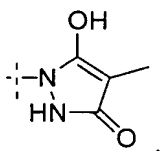
- COR<sup>2</sup>,
- CONR<sup>1</sup>S(O)<sub>2</sub>R<sup>9</sup>,
- COCH<sub>2</sub>SO<sub>2</sub>-thiazolyl optionally substituted with alkyl or amino,
- halo,
- NO<sub>2</sub>,
- OR<sup>1</sup>,
- R<sup>1</sup>,
- SR<sup>1</sup>,
- O-C<sub>1</sub>-C<sub>6</sub>-alkyl substituted by C<sub>3</sub>-C<sub>6</sub>-cycloalkyl,
- O-phenyl optionally substituted by SO<sub>2</sub>CH<sub>3</sub>,
- SO<sub>2</sub>NH<sub>2</sub>,
- SO<sub>2</sub>NR<sup>1</sup>R<sup>7</sup>,
- NR<sup>1</sup>R<sup>1</sup>,

•  $\text{NR}^1\text{COC}_1\text{-C}_6\text{alkyl}$ ,

•



•



5

•  $\text{C}_1\text{-C}_{10}\text{COR}^2$ ,

• phenyl optionally substituted with halo,  $\text{C}_1\text{-C}_4$  alkyl, or  $\text{C}_1\text{-C}_4$  alkoxy,

• tetrazolo;

10

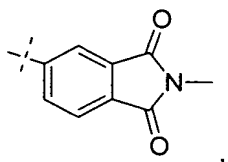
$\text{R}^7$  is

• phenyl or heteroaryl containing 3-6 C and 1-3 O, N, or S atoms, each optionally substituted by  $\text{C}_1\text{-C}_4$  alkyl, CN,  $\text{NO}_2$ ,  $\text{CO-C}_1\text{-C}_4\text{alkyl}$ ,  $\text{C}_1\text{-C}_4$  alkoxy, or  $\text{C}_1\text{-C}_4$  haloalkyl,

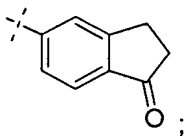
15

•  $\text{CO-R}^8$ ,

•



•



20

$\text{R}^8$  is

•  $\text{C}_1\text{-C}_6$  alkyl optionally substituted with  $\text{C}_1\text{-C}_4$  alkoxy,  $\text{N}(\text{CH}_3)_2$ , or one or two  $\text{CF}_3$ ,

•  $\text{C}_3\text{-C}_6\text{-cycloalkyl}$ ,

• phenyl optionally substituted with  $\text{C}_1\text{-C}_4$  alkoxy, halo, or  $\text{C}_1\text{-C}_4$  alkyl,

25

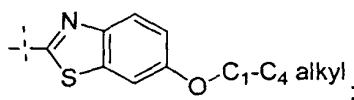
•  $\text{NH-phenyl}$  optionally substituted with  $\text{C}_1\text{-C}_4$  alkyl, halo,  $\text{C}_1\text{-C}_4$  alkoxy, or

C<sub>1</sub>-C<sub>4</sub> haloalkoxy,

- NH-cyclohexyl;

R<sup>9</sup> is

- C<sub>3</sub>-C<sub>6</sub> cycloalkyl,
- thienyl optionally substituted with C<sub>1</sub>-C<sub>4</sub> alkyl or isoxazolyl,
- pyridyl optionally substituted with -SO<sub>2</sub>-C<sub>1</sub>-C<sub>4</sub>alkyl,
- pyrazolyl optionally substituted with halo or C<sub>1</sub>-C<sub>4</sub> alkyl,
- isoxazolyl optionally substituted with C<sub>1</sub>-C<sub>4</sub> alkyl, or
- 



a is 0, 1, 2, 3, 4, or 5;

b is 0, 1, or 2;

d is 1, 2, or 3;

e is 1 or 2;

and pharmaceutically acceptable salts and esters thereof.

2. The compound of claim 1 wherein Y is

- halo,
  - R<sup>6</sup>,
  - SR<sup>1</sup>,
  - S(O)<sub>b</sub>-phenyl-CO<sub>2</sub>R<sup>1</sup>,
  - phenyl optionally fused to one or two phenyl rings, or to a 5- or 6-membered heterocycle containing one or more heteroatoms each independently selected from N, S, and O, or
  - a 5- or 6-membered heterocycle containing one or more heteroatoms each independently selected from N, S and O, optionally fused to a phenyl ring,
- each cyclic moiety being optionally substituted with one or more substituents independently selected from
- COR<sup>2</sup>,
  - halo,

- NO<sub>2</sub>,
- OR<sup>1</sup>,
- R<sup>1</sup>,
- SR<sup>1</sup>,
- 5      • SO<sub>2</sub>NR<sup>1</sup>R<sup>7</sup>,
- NR<sup>1</sup>R<sup>1</sup>,
- NR<sup>1</sup>COC<sub>1</sub>-C<sub>6</sub>alkyl,
- C<sub>1</sub>-C<sub>10</sub>COR<sup>2</sup>,
- phenyl,
- 10     • tetrazolo;

and pharmaceutically acceptable salts and esters thereof.

3. The compound of claim 1 wherein Y is
- 15      • phenyl optionally fused to one or two phenyl rings, or to a 5- or 6-membered heterocycle containing one or more heteroatoms each independently selected from N, S, and O, or
  - a 5- or 6- membered heterocycle containing one or more heteroatoms each independently selected from N, S and O, optionally fused to a phenyl ring,
  - 20     each cyclic moiety being optionally substituted with one or more substituents independently selected from
  - COR<sup>2</sup>,
  - halo,
  - NO<sub>2</sub>,
  - 25     • OR<sup>1</sup>,
  - R<sup>1</sup>,
  - SR<sup>1</sup>,
  - SO<sub>2</sub>NR<sup>1</sup>R<sup>7</sup>,
  - NR<sup>1</sup>R<sup>1</sup>,
  - 30     • NR<sup>1</sup>COC<sub>1</sub>-C<sub>6</sub>alkyl,
  - C<sub>1</sub>-C<sub>10</sub>COR<sup>2</sup>,
  - phenyl,
  - tetrazolo;

35      and d is 1 or 2;

and pharmaceutically acceptable salts and esters thereof.

4. The compound of claim 1 wherein

Y is

- 5
- phenyl optionally fused to one or two phenyl rings, or to a 5- or 6-membered heterocycle containing one or more heteroatoms each independently selected from N, S, and O, or
  - a 5- or 6- membered heterocycle containing one or more heteroatoms each independently selected from N, S and O, optionally fused to a phenyl ring,
- 10 each cyclic moiety being optionally substituted with one or more substituents independently selected from
- COR<sup>2</sup>,
  - halo,
  - NO<sub>2</sub>,
  - 15 • OR<sup>1</sup>,
  - R<sup>1</sup>,
  - SR<sup>1</sup>,
  - SO<sub>2</sub>NR<sup>1</sup>R<sup>7</sup>,
  - NR<sup>1</sup>R<sup>1</sup>,
  - 20 • C<sub>1</sub>-C<sub>10</sub>COR<sup>2</sup>,
  - phenyl,
  - tetrazolo;

Ar is

- 25
- phenyl optionally fused to a 5- or 6-membered heterocycle containing one or more heteroatoms each independently selected from O, S, and N, said bicyclic moiety being optionally fused to a phenyl, or
  - a 5- or 6-membered heterocycle containing one or more heteroatoms each independently selected from N, S, and O, optionally fused to phenyl;
- 30

and d is 1 or 2;

and pharmaceutically acceptable salts and esters thereof.

35



5. The compound of claim 1 wherein

Y is

- phenyl optionally fused to one or two phenyl rings, or to a 5- or 6-membered heterocycle containing one or more heteroatoms each independently selected from N, S, and O, or
- a 5- or 6- membered heterocycle containing one or more heteroatoms each independently selected from N, S and O, optionally fused to a phenyl ring, each cyclic moiety being optionally substituted with one or more substituents independently selected from

- COR<sup>2</sup>,
- halo,
- OR<sup>1</sup>,
- R<sup>1</sup>,
- NR<sup>1</sup>R<sup>1</sup>,

Ar is

- phenyl or
- a 5- or 6-membered heterocycle containing one or more N atoms;

a is 0, 1, 2, or 3; and

d is 1;

and pharmaceutically acceptable salts and esters thereof.

6. A compound selected from the group consisting of:

- 2-[4-(ethoxycarbonyl)phenoxy]-4-[(2R)-2-(((2R)-2-hydroxy-2-(3-pyridinyl)ethyl)amino)methyl)-3,4-dihydro-2H-chromen-6-yl]benzoic acid ;
- 4-[(2R)-2-(((2R)-2-hydroxy-2-(3-pyridinyl)ethyl)amino)methyl)-3,4-dihydro-2H-chromen-6-yl]-2-isobutylbenzoic acid;
- N-{3-[(2R)-2-(((2R)-2-hydroxy-2-(3-pyridinyl)ethyl)amino)methyl)-3,4-dihydro-2H-chromen-6-yl]benzoyl}-2-methylbenzenesulfonamide;
- 4-[(2R)-2-(((2R)-2-hydroxy-2-(3-pyridinyl)ethyl)amino)methyl)-3,4-dihydro-2H-chromen-6-yl]-2-isobutoxybenzoic acid;
- N-{3-[(2R)-2-(((2R)-2-hydroxy-2-(3-pyridinyl)ethyl)amino)methyl)-3,4-dihydro-2H-chromen-6-yl]benzoyl}-4-methoxybenzenesulfonamide;
- N-{3-[(2R)-2-(((2R)-2-hydroxy-2-(3-pyridinyl)ethyl)amino)methyl)-3,4-dihydro-2H-chromen-6-yl]benzoyl}-1-propanesulfonamide;

- 4-[(2R)-2-([[(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]-N-(4-methoxybenzoyl)benzenesulfonamide;
- N-(2-cyano-4-nitrophenyl)-3-[(2R)-2-([[(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]benzenesulfonamide;
- 5 2-(4-chlorophenoxy)-4-[(2R)-2-([[(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]benzoic acid;
- N-(4,6-dimethoxy-2-pyrimidinyl)-4-[(2R)-2-([[(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]-2-(trifluoromethoxy)benzenesulfonamide;
- 10 2-(4-fluorophenoxy)-4-[(2R)-2-([[(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]benzoic acid;
- 4-[(2R)-2-([[(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]-N-(3-methoxybenzoyl)benzenesulfonamide;
- 4-fluoro-N-[3-[(2R)-2-([[(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]benzoyl]benzenesulfonamide;
- 15 4-[(2R)-2-([[(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]-2-(4-methylphenoxy)benzoic acid;
- 4-[(2R)-2-([[(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]-2-(2-phenylethyl)benzoic acid;
- 20 3-chloro-4-[(2R)-2-([[(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]benzoic acid;
- N-(4-fluorobenzoyl)-4-[(2R)-2-([[(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]benzenesulfonamide;
- 4-[(2R)-2-([[(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]-3-methoxybenzoic acid;
- 25 4-[(2R)-2-([[(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]-2-phenoxybenzoic acid;
- N-(4-cyanophenyl)-4-[(2R)-2-([[(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]-2-(trifluoromethoxy)benzenesulfonamide;
- 30 4-[(2R)-2-([[(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]-N-(4-methoxy-6-methyl-2-pyrimidinyl)-2-(trifluoromethoxy)benzenesulfonamide;
- 4-[(2R)-2-([[(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]-N-(3,3,3-trifluoropropanoyl)benzenesulfonamide;

2-hydroxy-4-[(2R)-2-({[(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino}methyl)-3,4-dihydro-2H-chromen-6-yl]benzoic acid;

3-((1R)-2-({[(2R)-6-{4-[[[(4-fluorophenyl)amino]carbonyl]amino)sulfonyl]phenyl}-3,4-dihydro-2H-chromen-2-yl)methyl]amino}-1-hydroxyethyl)pyridine;

5 4-[(2R)-2-({[(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino}methyl)-3,4-dihydro-2H-chromen-6-yl]-N-(2-pyrimidinyl)benzenesulfonamide;

N-benzoyl-4-[(2R)-2-({[(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino}methyl)-3,4-dihydro-2H-chromen-6-yl]benzenesulfonamide;

10 4-[(2R)-2-({[(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino}methyl)-3,4-dihydro-2H-chromen-6-yl]-2-propoxybenzoic acid;

N-({4-[(2R)-2-({[(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino}methyl)-3,4-dihydro-2H-chromen-6-yl]-2-pyridinyl}carbonyl)-4-methoxybenzenesulfonamide;

3-((1R)-1-hydroxy-2-({[(2R)-6-{4-[[[(4-methylphenyl)amino]carbonyl]amino)sulfonyl]phenyl}-3,4-dihydro-2H-chromen-2-yl)methyl]amino}ethyl)pyridine;

15 3-((1R)-2-({[(2R)-6-{4-[[[(4-chloro-2-methylphenyl)amino]carbonyl]amino)sulfonyl]phenyl}-3,4-dihydro-2H-chromen-2-yl)methyl]amino}-1-hydroxyethyl)pyridine;

N-(ethoxyacetyl)-4-[(2R)-2-({[(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino}methyl)-3,4-dihydro-2H-chromen-6-yl]benzenesulfonamide;

20 N-(3,3-dimethylbutanoyl)-4-[(2R)-2-({[(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino}methyl)-3,4-dihydro-2H-chromen-6-yl]benzenesulfonamide;

4-[(2R)-2-({[(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino}methyl)-3,4-dihydro-2H-chromen-6-yl]-N-(4-methyl-2-pyrimidinyl)benzenesulfonamide;

25 4-[(2R)-2-({[(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino}methyl)-3,4-dihydro-2H-chromen-6-yl]-2-[4-(methylsulfonyl)phenoxy]benzoic acid;

4-[(2R)-2-({[(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino}methyl)-3,4-dihydro-2H-chromen-6-yl]-3-methylbenzoic acid;

4-{2-[(2R)-2-({[(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino}methyl)-3,4-dihydro-2H-chromen-6-yl]ethyl}benzoic acid;

30 N-(2,2-dimethylpropanoyl)-4-[(2R)-2-({[(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino}methyl)-3,4-dihydro-2H-chromen-6-yl]benzenesulfonamide;

3-[(1R)-2-({[(2R)-6-{4-[[[(anilino)carbonyl]amino)sulfonyl]phenyl}-3,4-dihydro-2H-chromen-2-yl)methyl]amino}-1-hydroxyethyl)pyridine;

- 2-ethoxy-4-[(2R)-2-([(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]benzoic acid;
- 4-[(2R)-2-([(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]-N-(4-methoxy-6-methyl-2-pyrimidinyl)benzenesulfonamide;
- 5 3-[(1R)-2-([(2R)-6-[4-([(cyclohexylamino)carbonyl]amino)sulfonyl]phenyl]-3,4-dihydro-2H-chromen-2-yl)methyl]amino)-1-hydroxyethyl]pyridine;
- N-(cyclopropylcarbonyl)-4-[(2R)-2-([(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]benzenesulfonamide;
- 10 2-chloro-5-fluoro-4-[(2R)-2-([(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]benzoic acid;
- 4-[(4-[R]-2-([(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino)methyl)-3,4-dihydro-2H-chromen-6-yl]benzoic acid;
- 4-[(2R)-2-([(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]-2-methylbenzoic acid;
- 15 2-fluoro-4-[(2R)-2-([(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]benzoic acid;
- 4-[(2R)-2-([(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]-3-propoxybenzoic acid;
- 4-[(2R)-2-([(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]-2-isopropoxybenzoic acid;
- 20 4-[(2R)-2-([(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]-N-(1,3-thiazol-2-yl)benzenesulfonamide;
- 4-[(2R)-2-([(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]-2-(4-methoxyphenoxy)benzoic acid;
- 25 3-(cyclopropylmethoxy)-4-[(2R)-2-([(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]benzoic acid;
- 4-[(2R)-2-([(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]benzenesulfonamide;
- 5-[(2R)-2-([(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]-4'-methyl-1,1'-biphenyl-2-carboxylic acid;
- 30 N-[6-[(2R)-2-([(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]-3-pyridinyl]benzenesulfonamide;
- 4-[(2R)-2-([(2R)-2-hydroxy-2-(3-pyridinyl)ethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]-N-(3-pyridinyl)benzenesulfonamide;

- 4-[(2R)-2-(((2R)-2-hydroxy-2-(3-pyridinyl)ethyl)amino)methyl)-3,4-dihydro-2H-chromen-6-yl]-2-methoxybenzoic acid;
- 4-chloro-N-{6-[(2R)-2-(((2R)-2-hydroxy-2-(3-pyridinyl)ethyl)amino)methyl)-3,4-dihydro-2H-chromen-6-yl]-3-pyridinyl}benzenesulfonamide;
- 5 4-[(2R)-2-(((2R)-2-hydroxy-2-(3-pyridinyl)ethyl)amino)methyl)-3,4-dihydro-2H-chromen-6-yl]-3-isobutoxybenzoic acid;
- N-{6-[(2R)-2-(((2R)-2-hydroxy-2-(3-pyridinyl)ethyl)amino)methyl)-3,4-dihydro-2H-chromen-6-yl]-3-pyridinyl}methanesulfonamide;
- 3-{2-[(2R)-2-(((2R)-2-hydroxy-2-(3-pyridinyl)ethyl)amino)methyl)-3,4-dihydro-2H-chromen-6-yl]ethyl}benzoic acid;
- 10 3-[(1E)-1-hexenyl]-4-[(2R)-2-(((2R)-2-hydroxy-2-(3-pyridinyl)ethyl)amino)methyl)-3,4-dihydro-2H-chromen-6-yl]benzoic acid;
- 3-[(2R)-2-(((2R)-2-hydroxy-2-(3-pyridinyl)ethyl)amino)methyl)-3,4-dihydro-2H-chromen-6-yl]-N-(2-pyrimidinyl)benzenesulfonamide;
- 15 4-[(2R)-2-(((2R)-2-hydroxy-2-(3-pyridinyl)ethyl)amino)methyl)-3,4-dihydro-2H-chromen-6-yl]-2-(2-methoxyethoxy)benzoic acid;
- 4-[(2R)-2-(((2R)-2-hydroxy-2-(3-pyridinyl)ethyl)amino)methyl)-3,4-dihydro-2H-chromen-6-yl]-2,6-dimethylbenzoic acid;
- 4-[(2R)-2-(((2R)-2-(6-amino-3-pyridinyl)-2-hydroxyethyl)amino)methyl)-3,4-dihydro-2H-chromen-6-yl]benzoic acid;
- 20 3-[(2R)-2-(((2R)-2-(6-amino-3-pyridinyl)-2-hydroxyethyl)amino)methyl)-3,4-dihydro-2H-chromen-6-yl]benzoic acid;
- (1R)-1-(6-amino-3-pyridinyl)-2-(((2R)-6-[4-(1H-tetraazol-5-yl)phenyl]-3,4-dihydro-2H-chromen-2-yl)methyl)amino]ethanol;
- 25 5-{4-[(2R)-2-(((2R)-2-hydroxy-2-(3-pyridinyl)ethyl)amino)methyl)-3,4-dihydro-2H-chromen-6-yl]phenyl}-3-phenyl-1,2,5,3,4-thiadiazole-2-carboxylic acid;
- 5-{4-[(2R)-2-(((2R)-2-hydroxy-2-(3-pyridinyl)ethyl)amino)methyl)-3,4-dihydro-2H-chromen-6-yl]phenyl}-2-furoic acid;
- 5-{4-[(2R)-2-(((2R)-2-hydroxy-2-(3-pyridinyl)ethyl)amino)methyl)-3,4-dihydro-2H-chromen-6-yl]phenyl}-2-thiophenecarboxylic acid;
- 30 5-{4-[(2R)-2-(((2R)-2-hydroxy-2-(3-pyridinyl)ethyl)amino)methyl)-3,4-dihydro-2H-chromen-6-yl]phenyl}-3-thiophenecarboxylic acid;
- 4-{4-[(2R)-2-(((2R)-2-hydroxy-2-(3-pyridinyl)ethyl)amino)methyl)-3,4-dihydro-2H-chromen-6-yl]phenyl}-2-thiophenecarboxylic acid;

- 6-[(2R)-2-([[(2R)-2-(6-amino-3-pyridinyl)-2-hydroxyethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]nicotinic acid;  
 5-[(2R)-2-([[(2R)-2-(6-amino-3-pyridinyl)-2-hydroxyethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]nicotinic acid;  
 5 2-[(2R)-2-([[(2R)-2-(6-amino-3-pyridinyl)-2-hydroxyethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]-4-pyridinecarboxylic acid;  
 1-([[(2R)-2-([[(2R)-2-(6-amino-3-pyridinyl)-2-hydroxyethyl]amino)methyl]-3,4-dihydro-2H-chromen-6-yl]carbonyl]amino)cyclopropanecarboxylic acid; and  
 4-[(2R)-2-([[(2R)-2-(3-chlorophenyl)-2-hydroxyethyl]amino)methyl]-3,4-dihydro-2H-  
 10 chromen-6-yl]benzoic acid (Example 344).
7. A method of treating a beta-3 adrenergic receptor-mediated condition comprising the step of administering to a patient in need thereof a pharmaceutically effective amount of a compound of claim 1.
- 15 8. A method of treating obesity comprising the step of administering to a patient in need thereof a pharmaceutically effective amount of a compound of claim 1.
9. A method of treating diabetes comprising the step of administering to a patient in need thereof a pharmaceutically effective amount of a compound of claim 1.
- 10 10. A method of treating a patient with impaired fasting glucose or impaired glucose tolerance comprising the step of administering to said patient in need thereof a pharmaceutically effective amount of a compound of claim 1.
11. A method of treating gastrointestinal disorders comprising the step of administering to a patient in need thereof a pharmaceutically effective amount of a compound of claim 1.
- 25 12. A method of treating hypertriglyceridemia, hypercholesteolemia, atherosclerotic disorders, or cardiovascular disorders comprising the step of administering to a patient in need thereof a pharmaceutically effective amount of a compound of claim 1.
- 30 13. A method for lowering high-density lipoprotein levels comprising the step of administering to a patient in need thereof a pharmaceutically effective amount of a compound of claim 1.

14. A method for treating urinary disorders comprising the step of administering to a patient in need thereof a pharmaceutically effective amount of a compound of claim 1.
- 5 15. The method of claim 14, wherein said urinary disorders is selected from the group consisting of pollakiuria and incontinence.
16. A method of treating a beta-3 adrenergic receptor-mediated condition comprising the step of administering to a patient in need thereof a pharmaceutically effective amount of a compound of claim 6.
- 10 17. A method of treating obesity comprising the step of administering to a patient in need thereof a pharmaceutically effective amount of a compound of claim 6.
18. A method of treating diabetes comprising the step of administering to a patient in need thereof a pharmaceutically effective amount of a compound of claim 6.
- 15 19. A method of treating a patient with impaired fasting glucose or impaired glucose tolerance comprising the step of administering to said patient in need thereof a pharmaceutically effective amount of a compound of claim 6.
- 20 20. A method of treating gastrointestinal disorders comprising the step of administering to a patient in need thereof a pharmaceutically effective amount of a compound of claim 6.
21. A method of treating hypertriglyceridemia, hypercholesteolemia, atherosclerotic disorders, or cardiovascular disorders comprising the step of administering to a patient in need thereof a pharmaceutically effective amount of a compound of claim 6.
- 25 22. A method for lowering high-density lipoprotein levels comprising the step of administering to a patient in need thereof a pharmaceutically effective amount of a compound of claim 6.
23. A method for treating urinary disorders comprising the step of administering to a patient in need thereof a pharmaceutically effective amount of a compound of claim 6.
24. The method of claim 23, wherein said urinary disorders is selected from the group



consisting of pollakiuria and incontinence.

- 5           25. A pharmaceutical composition comprising an effective amount of a compound of claim 1 or a pharmaceutically acceptable salt and esters thereof in combination with a pharmaceutically acceptable carrier.
- 10           26. A pharmaceutical composition for the treatment of obesity, diabetes, gastrointestinal disorders, hypertriglyceridaemia, hypercholesterolaemia, atherosclerosis, cardiovascular diseases, or urinary disorders comprising an effective amount of a compound of claim 1 or a pharmaceutically acceptable salt and ester thereof in combination with a pharmaceutically acceptable carrier.
- 15           27. A composition comprising an effective amount of a compound of claim 1 or a salt and esters thereof in combination with an inert carrier.
28. A pharmaceutical composition comprising an effective amount of a compound of claim 6 or a pharmaceutically acceptable salt and esters thereof in combination with a pharmaceutically acceptable carrier.
- 20           29. A pharmaceutical composition for the treatment of obesity, diabetes, gastrointestinal disorders, hypertriglyceridaemia, hypercholesterolaemia, atherosclerosis, cardiovascular diseases, or urinary disorders comprising an effective amount of a compound of claim 6 or a pharmaceutically acceptable salt and ester thereof in combination with a pharmaceutically acceptable carrier.
- 25           30. A composition comprising an effective amount of a compound of claim 6 or a salt and esters thereof in combination with an inert carrier.